Serial No.: 10/578,939 Filed: February 14, 2007

Page : 2 of 16

## Amendments to the Claims:

This listing of claims replaces all prior versions and listings of claims in the application:

## Listing of Claims:

- 1. (Currently Amended) An isolated detoxified pneumococcal neuraminidase or an antigenic portion thereof comprising an amino acid sequence of a non-detoxified neuraminidase with at least one amino acid substitution, wherein the detoxified pneumococcal neuraminidase or portion thereof has reduced activity as compared to the non-detoxified neuraminidase and wherein the detoxified pneumococcal neuraminidase or portion thereof is antigenic.
- 2. (Canceled)
- 3. (Canceled)
- 4. (Canceled)
- 5. (Canceled)
- 6. (Currently Amended) An The isolated detoxified pneumococcal neuraminidase or an antigenic portion thereof of claim 1, wherein the neuraminidase amino acid sequence comprises a deletion of at least 7% of the naturally occurring amino acids of the non-detoxified neuraminidase.
- 7. (Canceled)
- 8. (Canceled)
- 9. (Canceled)
- 10. (Canceled)
- 11. (Currently Amended) An The isolated detoxified pneumococcal neuraminidase or an antigenic portion thereof of claim 1, wherein the neuraminidase amino acid sequence comprises a deletion of at least 5 N-terminal amino acids of the non-detoxified neuraminidase.
- 12. (Currently Amended) An <u>The</u> isolated detoxified pneumococcal neuraminidase or an antigenic portion thereof of claim 1, wherein the neuraminidase amino acid sequence

Serial No.: 10/578,939 Filed: February 14, 2007

Page : 3 of 16

comprises a deletion of at least 10 N-terminal amino acids of <u>the</u> non-detoxified neuraminidase.

- 13. (Currently Amended) An The isolated detoxified pneumococcal neuraminidase or an antigenic portion thereof of claim 1, wherein the neuraminidase amino acid sequence comprises a deletion of at least 15 N-terminal amino acids of the non-detoxified neuraminidase.
- 14. (Currently Amended) An The isolated detoxified pneumococcal neuraminidase or an antigenic portion thereof of claim 1, wherein the neuraminidase amino acid sequence comprises a deletion of at least 10% of the C-terminal amino acids of the non-detoxified neuraminidase.
- 15. (Currently Amended) An The isolated detoxified pneumococcal neuraminidase or an antigenic portion thereof of claim 1, wherein the neuraminidase amino acid sequence comprises a deletion of at least 20% of the C-terminal amino acids of the non-detoxified neuraminidase.
- 16. (Currently Amended) An <u>The</u> isolated detoxified pneumococcal neuraminidase or an <u>antigenie</u> portion thereof <u>of claim 1</u>, wherein the <u>neuraminidase amino acid sequence</u> comprises a deletion of at least 30% of the C-terminal amino acids of <u>the</u> non-detoxified neuraminidase.
- 17. (Currently Amended) An The isolated detoxified pneumococcal neuraminidase or an antigenic portion thereof of claim 1, wherein the neuraminidase amino acid sequence comprises a deletion of at least 35% of the C-terminal amino acids of the non-detoxified neuraminidase.
- 18. (Currently Amended) A composition comprising an the isolated detoxified pneumococcal neuraminidase or an antigenic portion thereof of claim 1 and a pharmaceutically acceptable carrier.
- 19. (Original) The composition of claim 18, further comprising an adjuvant.
- 20. (Withdrawn) A method of generating antibodies specific to pneumococcal neuraminidase in a subject comprising administering to the subject a detoxified pneumococcal neuraminidase or an antigenic portion thereof.

Serial No.: 10/578,939 Filed: February 14, 2007

Page : 4 of 16

21. (Withdrawn) A method of reducing pneumococcal nasal carriage in a subject comprising administering to the subject a detoxified pneumococcal neuraminidase or an antigenic portion thereof.

- 22. (Withdrawn) A method of preventing pneumococcal infection in a subject comprising administering to the subject a detoxified pneumococcal neuraminidase or an antigenic portion thereof.
- 23. (Withdrawn) The method of claim 22, wherein the pneumococcal infection is meningitis.
- 24. (Withdrawn) The method of claim 22, wherein the pneumococcal infection is otitis media.
- 25. (Withdrawn) The method of claim 22, wherein the pneumococcal infection is pneumonia.
- 26. (Withdrawn) The method of claim 22, wherein the pneumococcal infection is hemolytic uremia.
- 27. (Withdrawn) A method of reducing or preventing pneumococcal nasal carriage in a subject comprising administering to the subject a detoxified pneumococcal neuraminidase or an antigenic fragment thereof under conditions that reduce or prevent the nasal carriage.
- 28. (Withdrawn) A method of reducing or preventing pneumococcal infection in a subject comprising administering to the subject a detoxified pneumococcal neuraminidase or an antigenic fragment thereof under conditions that reduce or prevent the infection.
- 29. (Withdrawn) The method of claim 28, wherein the pneumococcal infection is meningitis.
- 30. (Withdrawn) The method of claim 28, wherein the pneumococcal infection is otitis media.
- 31. (Withdrawn) The method of claim 28, wherein the pneumococcal infection is pneumonia.
- 32. (Withdrawn) The method of claim 28, wherein the pneumococcal infection is hemolytic uremia.
- 33. (Withdrawn) A method of reducing or preventing pneumococcal infection in a subject comprising administering to the subject a pneumococcal neuraminidase antibody or a fragment thereof under conditions that reduce or prevent the infection, wherein the administration step comprises contacting a mucosal surface of the subject with the antibody.

Serial No.: 10/578,939 Filed: February 14, 2007

Page : 5 of 16

34. (Withdrawn) The method of claim 33, wherein the pneumococcal infection is meningitis.

- 35. (Withdrawn) The method of claim 33, wherein the pneumococcal infection is otitis media.
- 36. (Withdrawn) The method of claim 33, wherein the pneumococcal infection is pneumonia.
- 37. (Withdrawn) The method of claim 33, wherein the pneumococcal infection is hemolytic uremia.
- 38. (Currently Amended) A composition comprising an the isolated detoxified pneumococcal neuraminidase or an antigenic portion thereof of claim 1 and a pharmaceutically acceptable carrier, wherein the composition is suitable for administration to a mucosal surface.
- 39. (Original) The composition of claim 38, wherein the composition is a nasal spray.
- 40. (Original) The composition of claim 38, wherein the composition is a nebulizer solution.
- 41. (Original) The composition of claim 38, wherein the composition is an aerosol inhalant.
- 42. (Original) A container comprising the composition of claim 38.
- 43. (Original) The container of claim 42, wherein the container is a nasal sprayer.
- 44. (Original) The container of claim 42, wherein the container is a nebulizer.
- 45. (Original) The container of claim 42, wherein the container is an inhaler.
- 46. (Withdrawn) A method of generating antibodies specific to pneumococcal neuraminidase in a subject comprising contacting the nasal mucosa of the subject with a detoxified pneumococcal neuraminidase or an antigenic portion thereof.
- 47. (Withdrawn) A method of reducing pneumococcal nasal carriage in a subject comprising contacting the nasal mucosa of the subject with a detoxified pneumococcal neuraminidase or an antigenic portion thereof.
- 48. (Withdrawn) A method of preventing pneumococcal infection in a subject comprising contacting a mucosal surface of the subject with a detoxified pneumococcal neuraminidase or an antigenic portion thereof.
- 49. (Withdrawn) The method of claim 48, wherein the pneumococcal infection is meningitis.
- 50. (Withdrawn) The method of claim 48, wherein the pneumococcal infection is otitis media.
- 51. (Withdrawn) The method of claim 48, wherein the pneumococcal infection is pneumonia.

Serial No.: 10/578,939 Filed: February 14, 2007

Page : 6 of 16

52. (Withdrawn) The method of claim 48, wherein the pneumococcal infection is hemolytic uremia.

- 53. (Withdrawn) A composition comprising a phosphocholine or an antigenic portion thereof of pneumococcal teichoic acid or pneumococcal lipoteichoic acid and a pharmaceutically acceptable carrier, wherein the composition is suitable for administration to a mucosal surface.
- 54. (Withdrawn) The composition of claim 53, wherein the composition is a nasal spray.
- 55. (Withdrawn) The composition of claim 53, wherein the composition is a nebulizer solution.
- 56. (Withdrawn) The composition of claim 53, wherein the composition is an aerosol inhalant.
- 57. (Withdrawn) A container comprising the composition of claim 53.
- 58. (Withdrawn) The container of claim 57, wherein the container is a nasal sprayer.
- 59. (Withdrawn) The container of claim 57, wherein the container is a nebulizer.
- 60. (Withdrawn) The container of claim 57, wherein the container is an inhaler.
- 61. (Withdrawn) A method of generating antibodies specific to pneumococcal neuraminidase in a subject comprising contacting the nasal mucosa of the subject with the composition of claim 53.
- 62. (Withdrawn) A method of reducing pneumococcal nasal carriage in a subject comprising contacting the nasal mucosa of the subject with the composition of claim 53.
- 63. (Withdrawn) A method of reducing or preventing pneumococcal infection in a subject comprising contacting a mucosal surface of the subject with the composition of claim 53.
- 64. (Withdrawn) The method of claim 63, wherein the pneumococcal infection is meningitis.
- 65. (Withdrawn) The method of claim 63, wherein the pneumococcal infection is otitis media.
- 66. (Withdrawn) The method of claim 63, wherein the pneumococcal infection is pneumonia.
- 67. (Withdrawn) The method of claim 63, wherein the pneumococcal is hemolytic uremia.
- 68. (Withdrawn) The composition of claim 38, further comprising a phosphocholine or an antigenic portion thereof of pneumococcal teichoic acid or pneumococcal lipoteichoic acid.

Serial No.: 10/578,939 Filed: February 14, 2007

Page : 7 of 16

69. (Withdrawn) The composition of claim 68, wherein the composition is a nasal spray.

- 70. (Withdrawn) The composition of claim 68, wherein the composition is a nebulizer solution.
- 71. (Withdrawn) The composition of claim 68, wherein the composition is an aerosol inhalant.
- 72. (Withdrawn) A container comprising the composition of claim 68.
- 73. (Withdrawn) The container of claim 72, wherein the container is a nasal sprayer.
- 74. (Withdrawn) The container of claim 72, wherein the container is a nebulizer.
- 75. (Withdrawn) The container of claim 72, wherein the container is an inhaler.
- 76. (Withdrawn) A method of generating antibodies specific to pneumococcal neuraminidase in a subject comprising contacting the nasal mucosa of the subject with the composition of claim 68.
- 77. (Withdrawn) A method of reducing pneumococcal nasal carriage in a subject comprising contacting the nasal mucosa of the subject with the composition of claim 68.
- 78. (Withdrawn) A method of preventing pneumococcal infection in a subject comprising contacting a mucosal surface of the subject with the composition of claim 68.
- 79. (Withdrawn) The method of claim 78, wherein the pneumococcal infection is meningitis.
- 80. (Withdrawn) The method of claim 78, wherein the pneumococcal infection is otitis media.
- 81. (Withdrawn) The method of claim 78, wherein the pneumococcal infection is pneumonia.
- 82. (Withdrawn) The method of claim 78, wherein the pneumococcal infection is hemolytic uremia.
- 83. (Withdrawn) A composition comprising a non-phosphocholine antigenic portion of pneumococcal teichoic acid or pneumococcal lipoteichoic acid and a pharmaceutically acceptable carrier, wherein the composition is suitable for administration to a mucosal surface.
- 84. (Withdrawn) The composition of claim 83, wherein the composition is a nasal spray.
- 85. (Withdrawn) The composition of claim 83, wherein the composition is a nebulizer solution.

Serial No.: 10/578,939 Filed: February 14, 2007

Page : 8 of 16

86. (Withdrawn) The composition of claim 83, wherein the composition is an aerosol inhalant.

- 87. (Withdrawn) A container comprising the composition of claim 83.
- 88. (Withdrawn) The container of claim 87, wherein the container is a nasal sprayer.
- 89. (Withdrawn) The container of claim 87, wherein the container is a nebulizer.
- 90. (Withdrawn) The container of claim 87, wherein the container is an inhaler.
- 91. (Withdrawn) A method of generating antibodies specific to pneumococcal neuraminidase in a subject comprising contacting the nasal mucosa of the subject with the composition of claim 83.
- 92. (Withdrawn) A method of reducing pneumococcal nasal carriage in a subject comprising contacting the nasal mucosa of the subject with the composition of claim 83.
- 93. (Withdrawn) A method of preventing pneumococcal infection in a subject comprising contacting a mucosal surface of the subject with the composition of claim 83.
- 94. (Withdrawn) The method of claim 93, wherein the pneumococcal infection is meningitis.
- 95. (Withdrawn) The method of claim 93, wherein the pneumococcal infection is otitis media.
- 96. (Withdrawn) The method of claim 93, wherein the pneumococcal infection is pneumonia.
- 97. (Withdrawn) The method of claim 93, wherein the pneumococcal infection is hemolytic uremia.
- 98. (Withdrawn) The composition of claim 38, further comprising a non-phosphocholine antigenic portion of pneumococcal teichoic acid or pneumococcal lipoteichoic acid.
- 99. (Withdrawn) The composition of claim 98, wherein the composition is a nasal spray.
- 100. (Withdrawn) The composition of claim 98, wherein the composition is a nebulizer solution.
- 101. (Withdrawn) The composition of claim 98, wherein the composition is an aerosol inhalant.
- 102. (Withdrawn) A container comprising the composition of claim 98.
- 103. (Withdrawn) The container of claim 102, wherein the container is a nasal sprayer.
- 104. (Withdrawn) The container of claim 102, wherein the container is a nebulizer.
- 105. (Withdrawn) The container of claim 102, wherein the container is an inhaler.

Serial No.: 10/578,939 Filed: February 14, 2007

Page : 9 of 16

106. (Withdrawn) A method of generating antibodies specific to pneumococcal neuraminidase in a subject comprising contacting the nasal mucosa of the subject with the composition of claim 98.

- 107. (Withdrawn) A method of reducing pneumococcal nasal carriage in a subject comprising contacting the nasal mucosa of the subject with the composition of claim 98.
- 108. (Withdrawn) A method of preventing pneumococcal infection in a subject comprising contacting a mucosal surface of the subject with the composition of claim 98.
- 109. (Withdrawn) The method of claim 108, wherein the pneumococcal infection is meningitis.
- 110. (Withdrawn) The method of claim 108, wherein the pneumococcal infection is otitis media.
- 111. (Withdrawn) The method of claim 108, wherein the pneumococcal infection is pneumonia.
- 112. (Withdrawn) The method of claim 108, wherein the pneumococcal infection is hemolytic uremia.
- 113. (Withdrawn) A composition comprising a phosphocholine antibody or a fragment thereof and a pharmaceutically acceptable carrier.
- 114. (Withdrawn) The composition of claim 113, wherein the composition is suitable for administration to a mucosal surface.
- 115. (Withdrawn) The composition of claim 113, wherein the composition is a nasal spray.
- 116. (Withdrawn) The composition of claim 113, wherein the composition is a nebulizer solution.
- 117. (Withdrawn) The composition of claim 113, wherein the composition is an aerosol inhalant.
- 118. (Withdrawn) A container comprising the composition of claim 113.
- 119. (Withdrawn) The container of claim 118, wherein the container is a nasal sprayer.
- 120. (Withdrawn) The container of claim 118, wherein the container is a nebulizer.
- 121. (Withdrawn) The container of claim 118, wherein the container is an inhaler.
- 122. (Withdrawn) A method of reducing pneumococcal nasal carriage in a subject comprising administering to the subject a phosphocholine antibody or a fragment thereof.

Serial No.: 10/578,939 Filed: February 14, 2007

Page : 10 of 16

123. (Withdrawn) The method of claim 122, wherein the administration comprising contacting the nasal mucosa of the subject with the antibody or fragment thereof.

- 124. (Withdrawn) A method of preventing pneumococcal infection in a subject comprising administering to the subject a phosphocholine antibody or a fragment thereof.
- 125. (Withdrawn) The method of claim 124, wherein the administration comprises contacting the nasal mucosa of the subject with the antibody or fragment thereof.
- 126. (New) The isolated detoxified pneumococcal neuraminidase or portion thereof of claim 1, wherein the amino acid sequence of the detoxified pneumococcal neuraminidase or portion thereof is at least 70% identical to the non-detoxified neuraminidase.
- 127. (New) The isolated detoxified pneumococcal neuraminidase or portion thereof of claim 1, wherein the amino acid sequence of the detoxified pneumococcal neuraminidase or portion thereof is at least 80% identical to the non-detoxified neuraminidase.
- 128. (New) The isolated detoxified pneumococcal neuraminidase or portion thereof of claim 1, wherein the amino acid sequence of the detoxified pneumococcal neuraminidase or portion thereof is at least 90% identical to the non-detoxified neuraminidase.